This Page Is Inserted by IFW Operations and is not a part of the Official Record

BEST AVAILABLE IMAGES

Defective images within this document are accurate representations of the original documents submitted by the applicant.

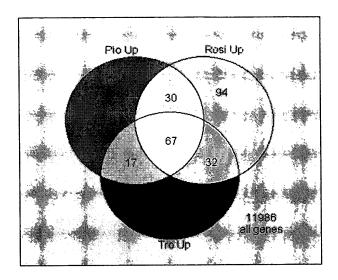
Defects in the images may include (but are not limited to):

- BLACK BORDERS
- TEXT CUT OFF AT TOP, BOTTOM OR SIDES
- FADED TEXT
- ILLEGIBLE TEXT
- SKEWED/SLANTED IMAGES
- COLORED PHOTOS
- BLACK OR VERY BLACK AND WHITE DARK PHOTOS
- GRAY SCALE DOCUMENTS

IMAGES ARE BEST AVAILABLE COPY.

As rescanning documents will not correct images, please do not report the images to the Image Problem Mailbox.

Figure 1: Venn overlap of genes up-regulated in response to PPARγ ligand treatment



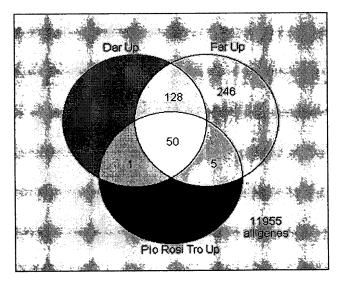


Figure 2: Venn overlap of genes down-regulated in response to PPAR γ ligand treatment

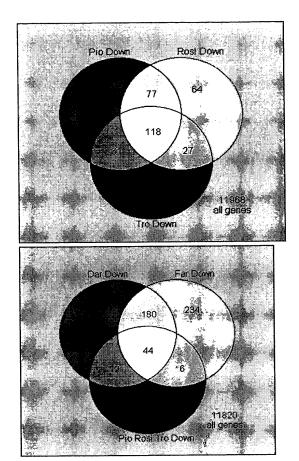


Figure 3. Fold change values for selected genes from core set of genes determined by Venn overlap to be 1.5 fold up or down-regulated by 24 hour Farglitazar, Darglitazone, Rosiglitazone, Pioglitazone, and Troglitazone treatment in 3T3-L1 adipocytes.

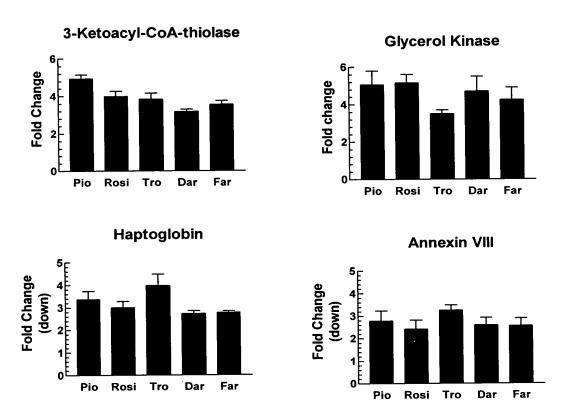
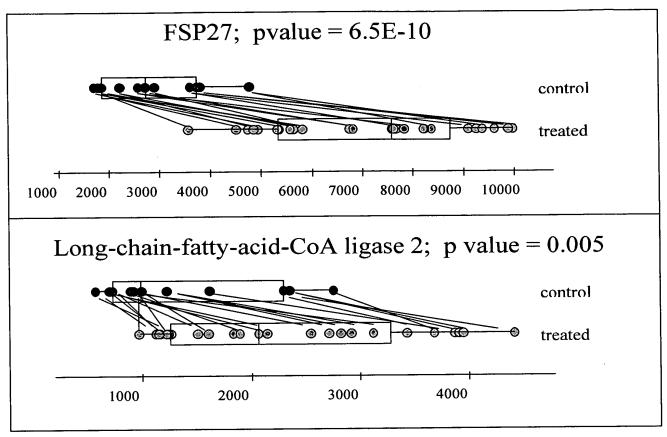
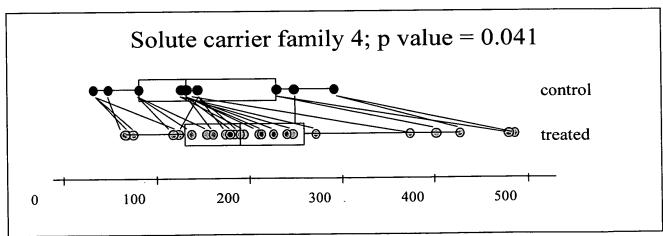
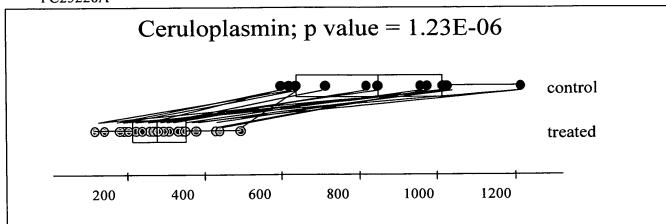
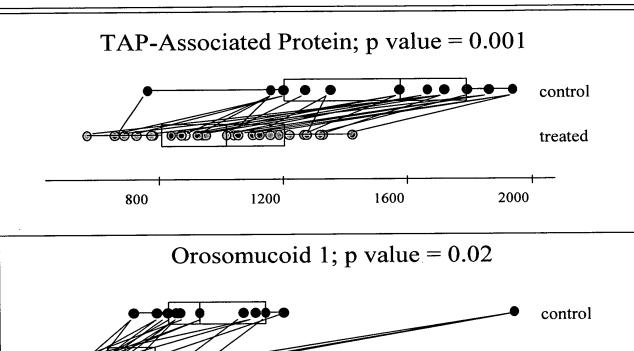


Figure 4. Box and whisker plots of selected genes up and down-regulated by PPAR γ ligand treatment.









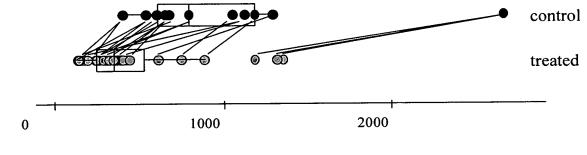
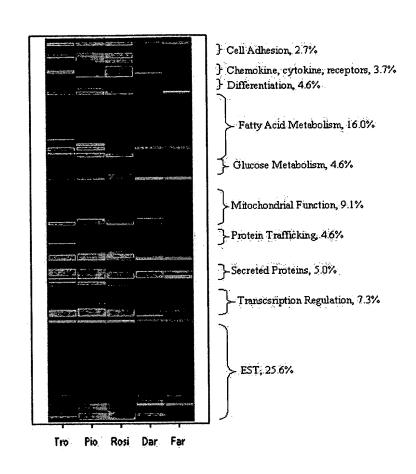


Figure 5. Heat map diagram of genes found to be 1.5 fold up or down-regulated by 24 hour PPAR $_{\gamma}$ ligand treatment in 3T3-L1 adipocytes. Diagram is colored by expression level and genes are grouped by biological function/pathway. Red represents up-regulated genes, black represents unchanged while green represents down-regulated genes



Log₂ Fold Change



Figure 6. PPAR $\gamma 1$ and PPAR $\gamma 2$ isoform RNA levels by Taqman analysis.

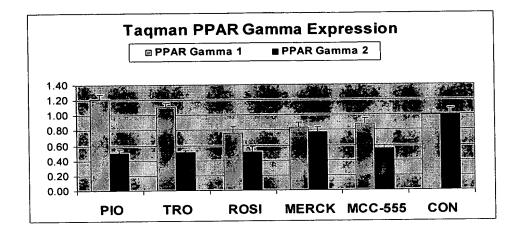


Figure 7. PPAR γ RNA levels by Affymetrix gene chip analysis.

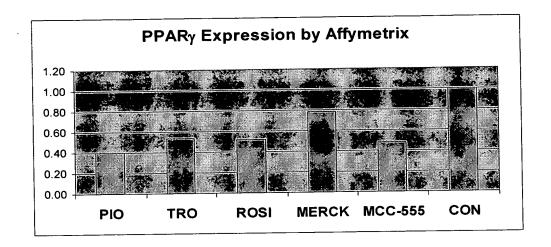
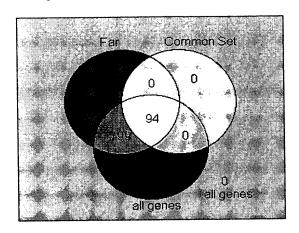
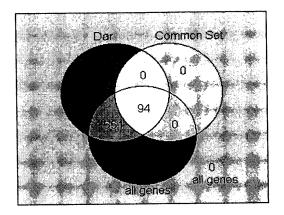
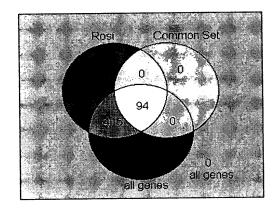


Figure 8. Venn diagram overlap of all genes up or down-regulated by the indicated PPAR γ ligands relative to the identified core set of genes.







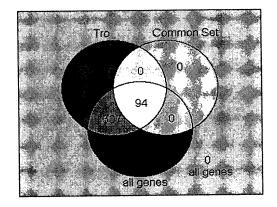


Figure 9. Ratio of total number of genes up or down-regulated by each PPAR γ ligand relative to the number of genes up or down-regulated by all ligands tested. This ratio is found to be correlated with the observed edema frequency of the particular ligands.

Edema Frequency vs. Off-target/On-target

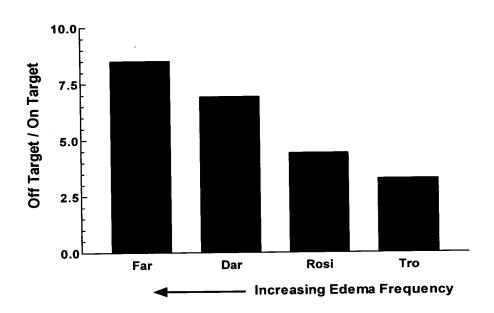


Figure 10. Venn diagram overlap of genes 1.5 Fold up and down-regulated by Rosiglitazone, Pioglitazone and Troglitazone treatment in experiment 2.

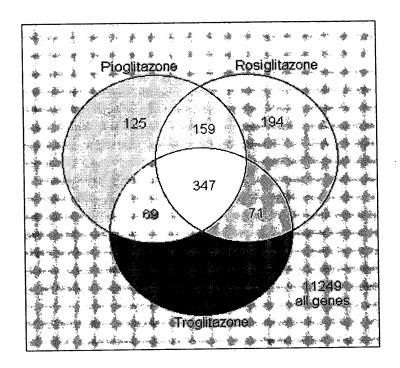


Figure 11. Venn diagram overlap of the core list of genes up and down-regulated by Farglitazar, Darglitazone, Rosiglitazone, Pioglitazone, and Troglitazone treatment, and the independently derived list of genes 1.5 fold up or down-regulated by Rosiglitazone, Pioglitazone and Troglitazone treatment in experiment 2.

